PCT

09/311,792

WORLD INTELLECTUAL PROPERTY ORGANIZATION



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁷ :		(11) International Publication Numbe	r: WO 00/70645
Н01Ј 35/06	A1	(43) International Publication Date:	23 November 2000 (23.11.00)

US

(21) International Application Number: PCT/US00/12920

(22) International Filing Date: 11 May 2000 (11.05.00)

(30) Priority Data:

13 May 1999 (13.05.99)

(71) Applicant: PHOTOELECTRON CORPORATION [US/US]; 5
Forbes Road – Suite 2, Lexington, MA 02421–7305 (US).

(72) Inventor: DINSMORE, Mark; 85 Horse Pond Road, Sudbury, MA 01776 (US).

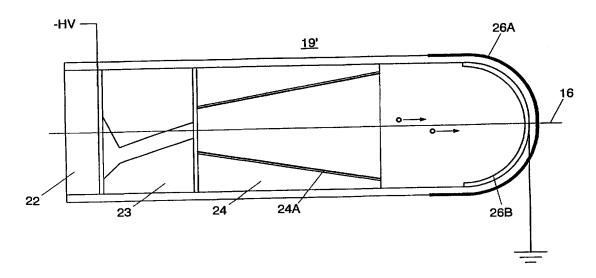
(74) Agent: LAPPIN, Mark, G.; McDermott, Will & Emery, 28 State Street, Boston, MA 02109 (US).

(81) Designated States: AU, CA, CN, JP, KR, European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE).

Published

With international search report.

(54) Title: MINIATURE X-RAY SOURCE AND FLEXIBLE PROBE



(57) Abstract

The invention is directed to a miniature x-ray source that may be coupled to a catheter comprising a source of free electrons (22), an electron multiplier (23), an electron accelerator (24) that diverges in a direction away from the electron source and has an inner surface coated with a semiconductor material, an x-ray emitting target (26B) and an x-ray transmissive window (26A).

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
AU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	-		-
					Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav	TM	Turkmenistan
BF	Burkina Faso	GR	Greece		Republic of Macedonia	TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	ΙE	Ireland	MN	Mongolia	UA	Ukraine
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NO	Norway	$\mathbf{z}\mathbf{w}$	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's	NZ	New Zealand		
CM	Cameroon		Republic of Korea	\mathbf{PL}	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
CU	Cuba	KZ	Kazakstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LI	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

MINIATURE X-RAY SOURCE AND FLEXIBLE PROBE

Background of the Invention

The present invention relates to a highly miniaturized, low power, programmable radiation source for use in delivering predefined doses of radiation to a predefined region and more particularly to a miniaturized radiation source mounted in a flexible probe.

5

10

15

20

25

30

In the field of medicine, radiation is used for diagnostic, therapeutic and palliative treatment of patients. The conventional medical radiation sources used for these treatments include large fixed position machines as well as small, transportable radiation generating probes. The current state of the art treatment systems utilize computers to generate complex treatment plans.

These systems apply doses of radiation that are known to inhibit the growth of new tissue because the radiation affects dividing cells more than the mature cells found in non-growing tissue. Thus, the tissue in the site of an excised tumor can be treated to prevent the regrowth of cancerous tissue and the recurrence of cancer. Alternatively, radiation can be applied to other areas of the body to inhibit tissue growth, for example the growth of new blood vessels inside the eye that can cause macular degeneration.

Conventional radiation treatments systems, such as the LINAC used for medical treatment, utilize a high power remote radiation source and direct a beam of radiation at a target area, such as tumor inside the body of a patient. This type of treatment is referred to as teletherapy because the radiation source is located a predefined distance, approximately one meter, from the target. This treatment suffers from the disadvantage that tissue disposed between the radiation source and the target is exposed to radiation.

An alternative treatment system utilizing a point source of radiation is disclosed in U.S. Patent No. 5,153,900 issued to Nomikos et al., U.S. Patent No. 5,369,679 to Sliski et al., and U.S. Patent No. 5,422,926 to Smith et al., all owned by the assignee of the present application, all of which are hereby incorporated by reference. This system includes a miniaturized, insertable probe capable of producing low power radiation in predefined dose geometries disposed about a predetermined location. This type of treatment is referred to as brachytherapy because the source is located close to or in some cases within the area receiving treatment. One advantage of brachytherapy is that the radiation is applied primarily to treat a predefined tissue volume, without significantly affecting the tissue adjacent to the treated volume.

Typical radiation therapy treatment involves positioning the insertable probe into or adjacent to the tumor or the site where the tumor or a portion of the tumor was removed to treat the tissue adjacent the site with a "local boost" of radiation. In order to facilitate controlled treatment of the site, it is desirable to support the tissue portions to be treated at a predefined distances from the radiation source. Alternatively, where the treatment involves the treatment of surface tissue or the surface of an organ, it is desirable to control the shape of the surface as well as the shape of the radiation field applied to the surface.

5

10

15

20

25

30

The treatment can involve the application of radiation, either continuously or intermittently, over an extended period of time. Therefore, it is desirable that the insertable probe be adjustably supported in a compliant manner to accurately position the radiation source with respect to the treated site and accommodate normal minor movements of the patient, such as movements associated with breathing.

In many x-ray therapeutic procedures, x-ray probes of the type generally disclosed in U.S. Patent No. 5,153,900 incorporate a relatively rigid tube enclosing an electron beam directed to an x-ray emitting target at its distal end. For example, in treatment of brain tumors, an x-ray probe having a rigid tube is used with a stereotactic frame affixed to the patient's skull, where the tube is advanced into a biopsy hole to the tumor location, as disclosed in U.S. Patent No. 5,369,679. The rigidity of the tube is useful in ensuring that the x-ray emitting target is properly located. In other cases, it is desirable to have a flexible tube leading to the x-ray emitting target, for example, where it is desirable to pass the probe up the urethra to the bladder, for treatment of the bladder. Such a flexible probe is disclosed in U.S. Patent No. 5,248,658.

However, it has been difficult to effectively treat tissue using the flexible probe of the latter patent.

Accordingly, it is an object of the present invention to provide an improved system for delivering radiation to a localized area.

It is a further object of the present invention to provide an improved highly miniaturized radiation source with a flexible probe.

Summary of the Invention

The present invention is directed to a miniaturized radiation source at the end of a flexible probe or catheter. The flexible catheter extends along a probe axis between a proximal end and a distal end of the catheter. The radiation source, at the distal end of the catheter, includes a substantially rigid housing disposed about a substantially evacuated interior region extending

along a beam axis from an electron source at an input end of the housing to a radiation transmissive window at an output end of the housing. The housing also may include, depending on the current capability of the electron source's electron emitter, a channel electron multiplier adapted for receiving electrons from the electron source and for producing free electrons at an output end of the channel electron multiplier and an electron accelerator adapted for establishing a potential difference in the interior region of the housing whereby the free electrons produced at the output end of the channel electron multiplier are accelerated toward a target at or near the window. The target produces x-radiation in response to incident accelerated free electrons.

5

10

15

20

25

30

Preferably, the electron accelerator includes a surface disposed about the beam axis between the electron source and the target on a ceramic and preferably monolithic, substrate. In one embodiment, the surface bears a semiconductor coating. The surface may be substantially conical in shape wherein the distance from the beam axis increases as a function of the distance from the electron source. The electron source can be a photocathode illuminated by laser energy, a field emitter or a thermionic emitter. The target and outer surface of the probe is preferably maintained at ground potential to reduce the risk of shock.

Brief Description of the Drawings

The foregoing and other objects of this invention, the various features thereof, as well as the invention itself, may be more fully understood from the following description, when read together with the accompanying drawings in which:

FIGURES 1A and 1B are a diagrammatic perspective view and a diagrammatic detail view, respectively, of a low power radiation source embodying the present invention;

FIGURES 2A and 2B are a perspective view and a sectional view, respectively, of an alternate form embodying the present invention;

FIGURE 3 is a diagrammatic representation of a sheath adapted for use with the apparatus of FIGURE 1;

FIGURE 4 is a schematic block diagram of the embodiment of FIGURE 1;

FIGURE 5 is a diagrammatic view of a low power radiation treatment system having a flexible probe embodying the present invention; and

FIGURE 6 is a diagrammatic view of a low power radiation source embodying the present invention.

Detailed Description of the Preferred Embodiments

The present invention is directed to a miniature, low power radiation producing probe which can be used for diagnostic, therapeutic and palliative treatment of patients. The radiation source in accordance with the present invention can made smaller than conventional radiation sources. In addition, the radiation source can be disposed at the distal end of the tip of a flexible (or rigid) tube or catheter which can be inserted into the body. In one embodiment, only a single high voltage wire is necessary for operation. The target and the outer surface of the probe are set at the ground potential to reduce the shock hazard of the device.

5

10

15

20

25

30

FIGURE 1A shows an x-ray apparatus 10 embodying the present invention. Apparatus 10 includes a housing 12 and an elongated cylindrical probe 14 extending from housing 12 along a reference axis 16 to an x-ray source assembly 19. Preferably, the probe 14 is flexible, as described below, but in some cases may be rigid. The housing 12 encloses a high voltage power supply 12A, a battery 12B and a control system 12C. The x-ray source assembly 19 has an electron source (cathode) 22 located in the distal end of the probe 14. Electron source 22 may be located in close proximity to a channel electron multiplier (CEM) 23 which receives electrons from the electron source 22. An accelerator 24 is positioned between the CEM 23 and an x-ray emitting (in response to incident accelerated electrons) target 26. The target underlies on x-ray transmissive window 27. Probe 14 is integral with the housing 12 and extends toward the target 26. In various embodiments, the x-ray emitting tip may be selectively shielded to control the spatial distribution of x-rays. In addition, the accelerator 24 may be magnetically shielded to prevent external magnetic fields from deflecting the beam away from designed impact points on the target.

FIGURE 1B shows an x-ray source assembly 19' for generating x-rays embodying the present invention. That source 19' is adapted for placement at the end of a cylindrical element (flexible or rigid). In an alternate form, shown in Figure 2A and 2B, an x-ray source 19" is positioned within a compact housing 12, the latter device is suitable to applying x-radiation body surface tissue.

In the various forms of x-ray source assembly 19, the electron beam generator 22 may include a thermionic emitter (driven by a low voltage power source or laser) or a photocathode (irradiated by an LED or laser source) or a field emitter. A single high voltage power supply 12A can be used to power the electron source (thermionic emitter) 22, the CEM 23 and accelerator 24. The accelerator 24 establishes an acceleration potential difference between the CEM 23 and the target 26 which is at ground potential. The beam generation and acceleration

components can be adapted to establish a thin (e.g. 1 mm or less in diameter) electron beam within the assembly 19 along a nominally straight axis 16.

5

10

15

20

25

30

Preferably, the CEM 23 is constructed as is well known and the electron multiplication value is predetermined as function of the intended use of the radiation source. Preferably, a high voltage of 1 Kvolt is connected to input end of the CEM.

Preferably, the accelerator is constructed from a monolithic ceramic material and includes an interior channel formed in the shape of the surface of a cone, although other shapes may be used, for example parabolic. The accelerator is disposed between the CEM 23 and the target 26 along the axis of the electron beam trajectory whereby the distance of the surface from the beam increases as a function of the distance from the CEM 23. Preferably, the surface includes a semiconductive coating 24A which ensures that the voltage gradient in the accelerator is smooth and linear and helps to prevent breakdown which occurs when the electrons hit the walls of the accelerator. In the preferred embodiment, the accelerator is the type described in commonly owned U.S. Patent Application Serial No. 09/211,144 entitled ELECTRON BEAM MULTISTAGE ACCELERATOR, which is incorporated herein by reference.

In one form of the invention, the outer cylindrical portion of the x-ray source assembly 19 is a hollow evacuated cylinder made of a molybdenum-rhenium, (Mo-Re), molybdenum (Mo) or mu-metal body with an interior diameter of 2 mm, and an exterior diameter of 3 mm. Preferably, beryllium (Be) cap and having a distance from the electron source to the target is less than 2mm. The target assembly 26 includes an emission element consisting of a small beryllium (Be) target element 26A which is coated on the side exposed to the incident electron beam with a thin film or layer 26B of a high-Z element, such as tungsten (W), uranium (U) or gold (Au). By way of example, with electrons accelerated to 30 keV-, a 2.2 micron thick tungsten film absorbs substantially all the incident electrons, while transmitting approximately 95% of any 30 keV-. 88% of any 20 keV-, and 83% of any 10 keV-x-rays generated in that layer. In the preferred embodiment, the beryllium target element 26A is 0.5 mm thick with the result that 95% of the xrays generated in directions normal to and toward the substrate 26A, and having passed through the tungsten target, are then transmitted through the beryllium substrate and outward at the distal end of assembly 19. While the target element 26A shown in FIGURE 3B is in the form of a hemispherical layer, other shaped elements may be used, such as those having disk-like or conical shapes.

In some forms of the target, the window element 26A may include a multiple layer film 26B, where the differing layers may have different emission characteristics. By way of example,

5

10

15

20

25

30

the first layer may have an emission (vs. energy) peak at a relatively low energy, and the second (underlying) layer may have an emission (vs. energy) peak at a relatively high energy. With this form of the invention, a low energy electron beam may be used to generate x-rays in the first layer (to achieve a first radiation characteristic) and high energy electrons may be used to penetrate through to the underlying layer (to achieve a second radiation characteristic). As an example, a 0.5 mm wide electron beam is emitted at the cathode and accelerated to 30 keVthrough the anode, with 0.1 eV transverse electron energies, and arrives at the target assembly 26 downstream from the anode, with a beam diameter of less than 1 mm at the target assembly 26. X-rays are generated in the target assembly 26 in accordance with preselected beam voltage, current, and target element 26B composition. The x-rays thus generated pass through the beryllium target substrate 26A with minimized loss in energy. As an alternative to beryllium, the target substrate 26A may be made of carbon or other suitable material which permits x-rays to pass with a minimum loss of energy. An optimal material for target substrate 26A is carbon in its diamond form, since that material is an excellent heat conductor. Using these parameters, the resultant x-rays have sufficient energy to penetrate into soft tissues to a depth of a centimeter or more, the exact depth dependent upon the x-ray energy distribution.

The apparatus of FIGURES 2A and 2B is particularly adapted for full implantation into a patient, where the housing 12 a biocompatible outer surface and encloses both a high voltage power supply circuit 12A for establishing a drive voltage for the beam generator 22, and an associated battery 12B for driving that circuit 12A. In this case, an associated controller 12C establishes control of the output voltage of the high power supply circuit 12A, in the manner described below.

The apparatus of FIGURES 1A and 1B may be used in a manner where only the probe 14 and x-ray source assembly 19 are inserted into a patient while the housing 12 remains outside the patient, i.e., a transcutaneous form. In the latter form, some or all of the various elements shown within housing 12 may alternatively be remotely located.

In the transcutaneous form, the apparatus 10 may be used with an elongated closed end (or cup-shaped) sheath 34, as shown in FIGURE 3, having a biocompatible outer surface, for example, fabricated of medical grade aliphatic polyurethane, as manufactured under the trademark Tecoflex by Thermedics, Inc., Woburn, Massachusetts. With this configuration, the probe 14 is first inserted into the sheath 34. The sheath 34 and probe 14 are then inserted into the patient through the skin. Alternatively, a port may be inserted through the skin and attached to it,

as for example a Dermaport port manufactured by Thermedics Inc., Woburn, Massachusetts. The probe 14 is then inserted into the port.

The lining of the sheath or port can be configured as an x-ray shield by introducing barium sulfate or bismuth trioxide, or other x-ray shielding materials, into the sheath. If necessary, the probe 14 and housing 12 can be secured to the patient's body to prevent any relative motion during the extended time of treatment. An exemplary sheath 34 is shown in FIGURE 3.

5

10

15

20

25

30

In one embodiment of the apparatus as shown in FIGURES 1A and 1B, the main body of the probe 14 can be made of a magnetically shielding material such as a mu-metal.

Alternatively, the probe 14 can be made of a non-magnetic metal, preferably having relatively high values for Young's modulus and elastic limit. Examples of such material include molybdenum, rhenium or alloys of these materials. The outer cylindrical shell of the accelerator 24 can be made of the outer shell metal. The inner or outer surface of probe 14 can then be coated with a high permeability magnetic alloy such as permalloy (approximately 80% nickel and 20% iron), to provide magnetic shielding. Alternatively, a thin sleeve of mu-metal can be fitted over, or inside of that shell of accelerator 24. The x-ray apparatus 10 can then be used in environments in which there are dc and ac magnetic fields due to electrical power, the field of the earth, or other magnetized bodies nominally capable of deflecting the electron beam from the probe axis.

In implantable configurations, such as those of Figures 2A and 2B, the power supply 12A and target assembly 26 are preferably enclosed in a capsule to prevent current flow from the x-ray source to the patient. The closed housing 12 and probe 14 are, thus, encapsulated in a continuous outer shell of appropriate shielding material such as those mentioned previously.

The high voltage power supply 12A in each of the illustrated embodiments preferably satisfies three criteria: 1) small in size; 2) high efficiency to enable the use of battery power; and 3) independently variable x-ray tube voltage and current to enable the unit to be programmed for specific applications. A high-frequency, switch-mode power converter is used to meet these requirements. The most appropriate topology for generating low power and high voltage is a resonant voltage converter working in conjunction with a high voltage, Cockroft-Walton-type multiplier. Low-power dissipation, switch-mode power-supply controller-integrated circuits (IC) are currently available for controlling such topologies with few ancillary components.

The embodiment of FIGURES 2A and 2B can also be adapted for superficial usage, that is for direct placement on the skin of a patient. This form of the invention is particularly useful

for x-ray treatment of skin lesions or tumors, or other dermatological applications. In FIGURES 2A and 2B, elements that correspond to elements in the embodiment of FIGURES 1A and 1B are denoted with the same reference designations. Apparatus 10' generates an electron beam in a channel 40 enclosed within housing 12, where that channel 40 corresponds to probe 14. In the present embodiment, of FIGURES 2A and 2B, the x-ray source assembly 19 functions in a manner similar to that described above. With the configuration of FIGURES 2A and 2B, low power x-rays may be directed to a desired skin region of a patient.

5

10

15

20

25

30

In all of the above-described embodiments, the x-ray emission element of the target assembly is adapted to be adjacent to or within the region to be irradiated. The proximity of the emission element to the targeted region, e.g. the tumor, eliminates the need for the high voltages of presently used machines, to achieve satisfactory x-ray penetration through the body wall to the tumor site. The low voltage also concentrates the radiation in the targeted tumor, and limits the damage to surrounding tissue and surface skin at the point of penetration. For example, the delivery of 4000 rads, as is required after a mastectomy, with a 40 kV, 20 μ A electron beam, may require approximately 1 to 3 hours of radiation. However, since the x-ray source is, in this preferred embodiment, insertable proximate to, or into, the region-to-be-irradiated risk of incidental radiation exposure to other parts of the patient's body is significantly reduced.

Further, specificity in treating tumors may be achieved by tailoring the target and shield geometry and material at the emission site, for example as disclosed in U.S. Patent No. PHLL-111, assigned to the assignee of the present invention, . This tailoring facilitates the control of energy and the spatial profile of the x-ray emission to ensure more homogenous distribution of the radiation throughout the targeted tumor.

FIGURE 4 is a schematic representation of the x-ray source apparatus 10 shown in FIGURE 1A. In that preferred configuration, the housing 12 is divided into a first portion 12' and a second portion 12". Enclosed within the first housing portion 12' is a rechargeable battery 12B, a recharge network 12D for the battery 12B, which is adapted for use with an external charger 50, and a telemetry network 12E, adapted to be responsive to an external telemetry device 52 to function in the manner described below. That portion 12' is coupled by cables to the second housing portion 12". The second housing portion 12" includes the high voltage power supply 12A, controller 12C and the probe 14, as well as the electron beam generating element 22. In one embodiment, the electron beam generator includes a thermionic emitter 22 driven by the power supply 12A. In operation, power supply 12A heats the thermionic emitter 22, which in turn generates electrons which are then accelerated toward the anode 24. The anode 24 attracts

the electrons, but passes them through its central aperture toward the target assembly 26. The controller 12C controls the power supply 12A to dynamically adjust the cathode voltage, the electron beam current, and temporal parameters, or to provide pre-selected voltage, beam current, and temporal parameters.

Also illustrated, is an alternative electron beam generator which includes a photoemitter 22 irradiated by a light source 56, such as a diode laser or LED, powered by a driver 55. The light is focused on the photoemitter 22 by a focusing lens 58.

5

10

15

20

25

30

In the illustrated embodiment, external telemetry device 52 and telemetry network 12E cooperate to permit external control (dynamic or predetermined) control over the power supply 12A and temporal parameters. In embodiments when the housing 12" is not implanted, but where only probe 14 extends into a patient's body, the controller 12C may directly be used to control operation and in that case there is no need for network 12E.

FIGURES 5 and 6 show a diagrammatic view of radiation treatment apparatus 200 including a flexible probe 214. The apparatus 200 includes a high voltage source 218, a laser (or other optical) source 220, a probe assembly 214, and a radiation source assembly 226. According to one aspect of the invention, the apparatus 200 provides the required flexibility, without using strong magnetic fields, by locating electron source components 222, 223 and accelerator 224 near the target 228 in the distal end of the probe 214. The probe assembly 214 couples both the laser source 220 and the high voltage feed 218 to the radiation source assembly 226. Preferably, the probe assembly includes flexible fiber optical cable 202 enclosed in a small-diameter flexible metallic tube 204.

The radiation source assembly 226, which can be for example 1 to 2 cm in length, extends from the end of the probe assembly 214 and includes a shell which encloses the target 228. According to one embodiment, the radiation source assembly 226 is rigid in nature and generally cylindrical in shape. In this embodiment the cylindrical shell enclosing the radiations source assembly 226 can be considered to provide a housing for the electron beam source as well as a tubular probe extending from the housing along the electron beam path. The inner surface 226A of the assembly 226 is lined with an electrical insulator, while the external surface of the assembly 226 is electrically conductive. According to a preferred embodiment, the radiation source assembly is hermetically sealed to the end of the probe assembly 214, and evacuated. According to another embodiment, the entire probe assembly 214 is evacuated.

The terminal end 202A of the fiber optical cable 202 is preferably coated, over at least part of its area, with a semitransparent photoemissive substance such as, Ag-O-Cs, thus forming

a photocathode 222. A high voltage conductor 208, embedded in the fiber optical cable 202, conducts electrons to the cathode 222 (if necessary), the electron multiplier 223 and the accelerator 224 from the high voltage source 218. Similarly, the flexible tube 204 couples a ground return from the target 228 to the high voltage source 218, thereby establishing a high voltage field between the cathode 216 and the target 228. The fiber optical cable 202 acts as an insulating dielectric between the high voltage conductor 208 and the grounded flexible tube 204.

5

10

15

20

25

30

In order to eliminate scattering of the light in the fiber optic cable 202 by the high voltage wire 208, the fiber optic cable 202 can have an annular configuration. The light from the laser 220 travels down the annular core of the fiber optic cable 202. Cladding can be provided on each side of the core having an index of refraction so as to reflect the light beam incident on the interface back into the core. The grounded flexible metal tube 204 can surround the outer cladding.

As in previously described embodiments, the target 228 can be for example, beryllium, (Be), coated on one side with a thin film or layer 228A of a higher atomic number element, such as tungsten (W) or gold (Au).

In operation, the small semiconductor laser 220 shining down the fiber optical cable 202 activates the transmissive photocathode 222 which generates free electrons 216. The high voltage field between the cathode 222 and target 228 accelerates these electrons, thereby forcing them to strike the surface 228A of target 228 and produce x-rays. In order to generate, for example, 20 μ A of current from an Ag-O-Cs photocathode 222 with a laser 220 emitting light at a wavelength of 0.8 m, the 0.4% quantum efficiency of this photocathode 222 for this wavelength requires that the laser 220 emits 7.5 mW optical power. Such diode lasers are readily commercially available. According to the invention, the photoemissive surface which forms cathode 222 can, in fact, be quite small. For example, for a current density at the cathode 222 of 1 A/cm², the photoemitter's diameter need only be approximately 50 μ m.

One difficult fabrication aspect of this invention is the fabrication of the photocathode 222, which for practical substances, with reasonable quantum efficiencies above 10^{-3} , should be performed in a vacuum. This procedure can be carried out with the fiber optical cable 202 positioned in a bell jar, where for example, an Ag-O-Cs photosurface is fabricated in the conventional manner. Subsequently, without exposure to air, the optical cable 202 can be inserted into the tube 204. The end 202B can be vacuum sealed to the flexible tube 204.

In the above embodiments, the probe 14 or 214, along with its associated target 26, or 228, can be coated with a biocompatible outer layer, such as titanium nitride on a sublayer of nickel. For additional biocompatibility, a sheath of, for example, polyurethane can be fitted over the probe, such as that illustrated in FIGURE 3.

The invention may be embodied in other specific forms without departing from the spirit or essential characteristics thereof. The present embodiments are therefore to be considered in respects as illustrative and not restrictive, the scope of the invention being indicated by the appended claims rather than by the foregoing description, and all changes which come within the meaning and range of the equivalency of the claims are therefore intended to be embraced therein.

5

10

What is claimed is

1	1. A miniature radiation source comprising:
2	a flexible catheter extending along a probe axis between a proximal end and a distal end
3	of the catheter;
4	a radiation source having a substantially rigid housing defining a substantially evacuated
5	interior region extending along a beam axis between an electron source at an input end of the
6	housing and a radiation transmissive window at an output end of the housing, the housing having
7	a target responsive to incident accelerated free electrons to emit radiation whereby the radiation
8	emitted therefrom is directed through the radiation transmissive window, the housing having the
9	input end affixed to the distal end of the catheter;
10	means responsive to a signal at the proximal end of the catheter for selectively activating
11	the electron source to emit free electrons in the interior region
12	a channel electron multiplier adapted for receiving electrons from the electron source, at
13	an input end, and adapted for producing free electrons at an output end; and
14	an electron accelerator adapted for establishing a potential difference in the interior
15	region of the radiation source whereby the free electrons are accelerated toward the target.

- 1 2. A miniature radiation source according to claim 1, wherein the electron accelerator
- 2 includes a surface disposed about the beam axis between the electron source and the target and
- 3 being characterized as having an increasing distance from the beam axis as a function of
- 4 distance from the electron source, the surface bearing a semiconductor coating.
- 1 3. A miniature radiation source according to claim 2, wherein the surface is established by
- 2 a monolithic ceramic element.
- 1 4. A miniature radiation source according to claims 2, wherein the surface is a surface of
- 2 revolution.
- 1 5. A miniature radiation source according to claims 2, wherein the function is linear.

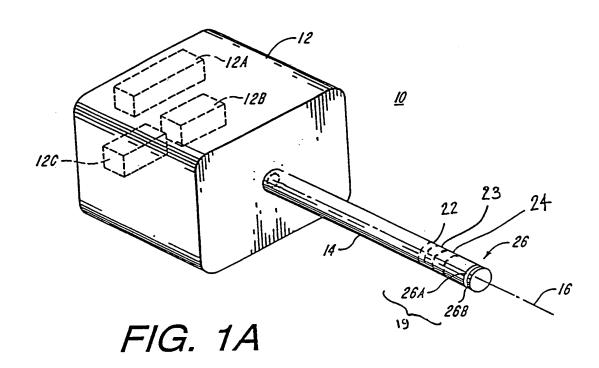
1 6. A miniature radiation source according to claim 1, wherein the electron source is

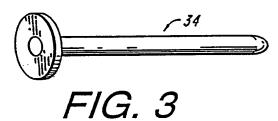
- 2 selected from the group including a photocathode illuminated by laser energy, a field emitter,
- 3 and a thermionic emitter.
- 1 7. A miniature radiation source according to claim 1, wherein the target is at ground
- 2 potential.
- 1 8. A therapeutic radiation source adapted for coupling to a catheter, comprising:
- a radiation source having a substantially rigid housing defining a substantially evacuated
- 3 interior region extending along a beam axis between an electron source at an input end of the
- 4 housing and a radiation transmissive window at an output end of the housing, the radiation
- 5 source having a target responsive to incident accelerated free electrons to emit radiation disposed
- 6 along the beam axis whereby the radiation emitted therefrom is directed through the radiation
- 7 transmissive window, the electron source generating the electrons in response to a signal
- 8 communicated through the catheter, the window being at ground potential; and
- a ceramic monolithic accelerator for accelerating the electrons along the beam axis, the
- 10 accelerator having a hollow interior for passing electrons therethrough, the interior diverging
- away from electron source, the interior being coated with a semiconductor coating to provide a
- 12 smooth voltage gradient along the beam axis and to reduce secondary emissions from the
- 13 accelerator.
- 1 9. A therapeutic radiation source according to claim 8, further comprising means
- 2 connecting the coating to a control voltage in the catheter for modifying electron acceleration
- 3 through the accelerator.
- 1 10. A therapeutic radiation source according to claim 8, further comprising a channel
- 2 electron multiplier adjacent to the source for multiplying electrons for acceleration within the
- 3 accelerator.

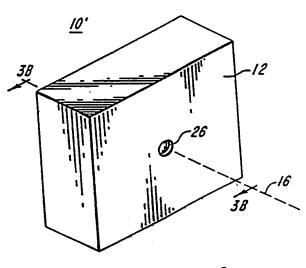
1 11. A therapeutic radiation source according to claim 8, wherein the interior comprises a

- 2 single cylindrical wall.
- 1 12. A therapeutic radiation source according to claim 8, wherein the electron source is
- 2 selected from the group consisting of a photocathode illuminated by laser energy, a field
- 3 emitter, and a thermionic emitter.
- 1 13. A therapeutic radiation source according to claim 8, wherein the source comprises a
- 2 channel electron multiplier.
- 1 14. A radiation source for attachment to a therapeutic probe, the radiation source having an
- 2 elongated rigid housing defining a substantially evacuated interior region extending along a
- 3 beam axis, said radiation source comprising:
- 4 an electron source for producing free electrons;
- a ceramic monolithic accelerator for accelerating the free electrons toward a radiation
- 6 target, the accelerator having a hollow interior for passing electrons therethrough, the interior
- 7 diverging away from electrons of increasing acceleration, the interior being coated with a
- 8 semiconductor coating to provide a substantially smooth voltage gradient between high voltage
- 9 and ground potential and to reduce secondary emissions from the accelerator, the radiation target
- 10 converting accelerated electrons into radiation;
- a radiation window, coupled to ground potential, for passing the radiation through the
- 12 housing.
- 1 15. A radiation source according to claim 14 where in said electron source comprises an
- 2 electron generator and an electron multiplier, said electron multiplier being adapted for
- 3 producing a quantity of free electrons as a function of a quantity of electrons produced by said
- 4 electron generator.

1/5









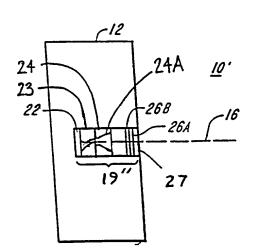
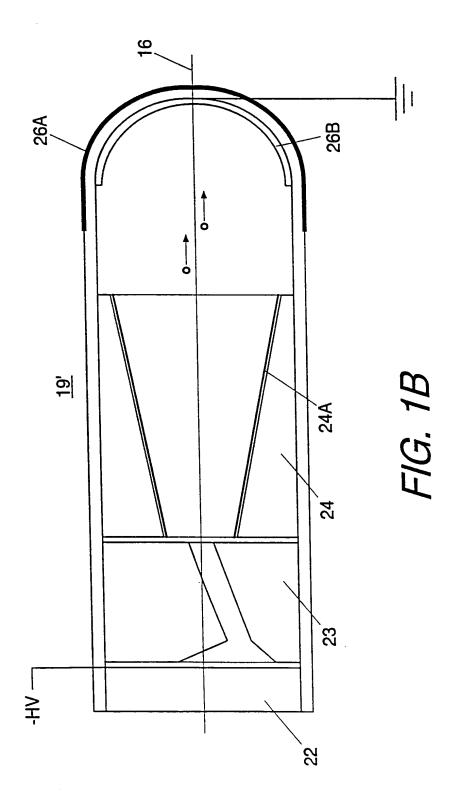
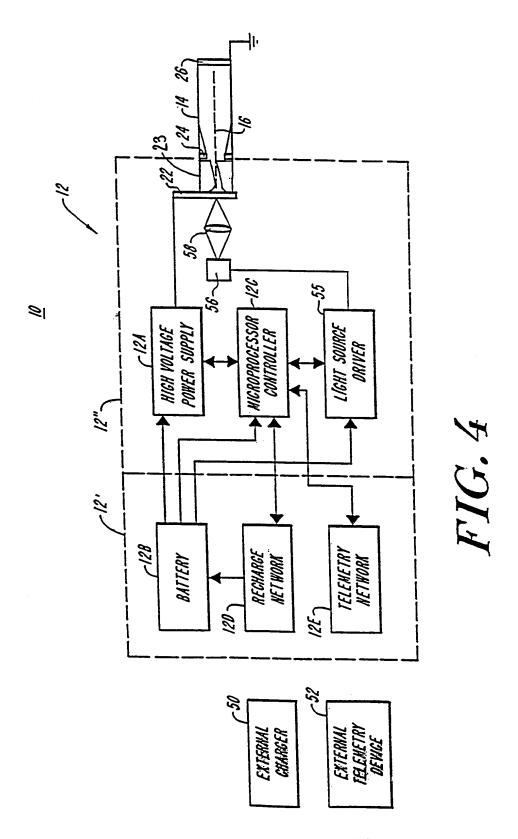
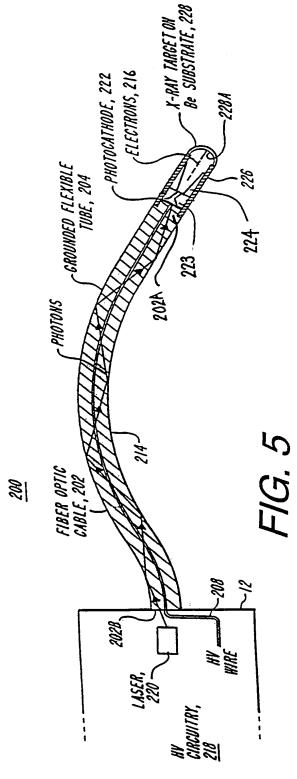


FIG. 2B





SUBSTITUTE SHEET (RULE 26)



SUBSTITUTE SHEET (RULE 26)

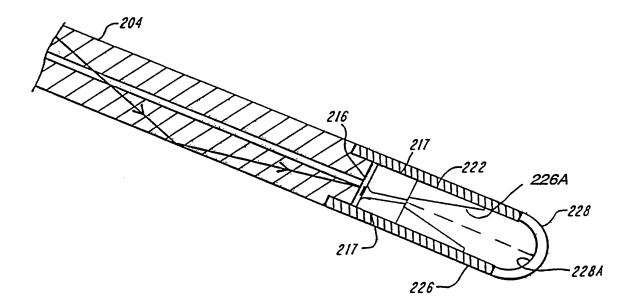


FIG. 6

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US00/12920

A. CLASSIFICATION OF SUBJECT MATTER IPC(7) :HOLD 35/66 US CL :378/136 According to international Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) U.S. : 378/136 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Catagory* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Y. US 5,153,900 A (Nomikos et al) 06 October 1992 (06/10/92) See the entire document. A. US 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) See the entire document. Discontinuous programs asses of the servicine of other continuation of Box C. See patent family annex. The document which may drow oblide an priority claim of the service of th			···					
According to Informational Patent Classification (IPC) or to both national classification and IPC B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) U.S.: 378/136 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Cutegory* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. U.S. 5, 153, 900 A (Nomikos et al) 06 October 1992 (06/10/92) See the entire document. 3. U.S. 5, 576, 549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) See the entire document. 4. Special categories of data documents. 4. Special categories of data documents. 5. Special categories of data documents. 5. Special categories of data documents. 6. See the entire document is described in the continuation of Box C. 5. Special categories of data documents. 6. See the entire documents. 7. Special categories of data documents. 7. Special categories of data documents. 7. Special categories of data documents. 8. Special categories of data documents. 8. Special categories of data documents. 8. See patent family annex. 8. See patent family annex. 8. See patent family annex. 9. See patent family annex. 9. See the entire documents are blisted in the continuation of Box C. 10. See patent family annex. 11. See patent family annex. 12. See patent family annex. 13. See patent family annex. 14. See the entire documents are blisted and tradection of a categories data on a categories of annex to exclusion categories annex to exclusion								
### FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) U.S.: 378/136 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Category* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. U.S. 5, 153, 900 A (Nomikos et al) 06 October 1992 (06/10/92) See the entire document. A. U.S. 5, 576, 549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) See the entire document. **See the entire document. **See the entire document. **See the entire documents are listed in the continuation of Box C. See patent family annex. **To document defining the general state of the air which is not considered to do for inclination of other viscosity of particular privateurs of the proprietation of particular privateurs of the proprietation and th	US CL	US CL :378/136						
Minimum documentation searched (classification system followed by classification symbols) U.S.: 378/136 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched like and the search of the search documents are included in the fields searched like search documents are included in the fields searched like search documents are listed in the continuation, where appropriate, of the relevant passages Relevant to claim No. Y. U.S. 5, 153, 900 A (Nomikos et al) 06 October 1992 (06/10/92) See the entire document. Y. See the entire document. **See the entire document of the search state of the art which is not considered to be of particular relevance to be of particular relevance to be of particular relevance to the search of the profits of an other of the search of another collision or other means **Committee of the search of the sea			national classification and IPC					
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Commentation searched other than minimum documentation to the extent that such documents are included in the fields searched that base consulted during the international search (name of data base and, where practicable, search terms used) Commentation of document, with indication, where appropriate, of the relevant passages Relevant to claim No.		····	1 by classification symbols)					
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) C. DOCUMENTS CONSIDERED TO BE RELEVANT Category* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. 1,6,7 2-5, 8-15 X US 5,153,900 A (Nomikos et al) 06 October 1992 (06/10/92) See the entire document. * US 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) See the entire document. * See the entire document. * See the entire document. * See the entire document when the store of the art which is not considered to be disputable relevant when the store doubts on growing claim() and which is growing the store of the store of another order or other means the store of another order or other special reason (as specified) To document referring to an oral disclosure, use, exhibitions or other special reason (as specified) To document referring to an oral disclosure, use, exhibitions or other means the parent referring to an oral disclosure, use, exhibitions or other means the promy data channel Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Tademarks Box PCT Vashington, D.C. 20231		•	-,,					
C. DOCUMENTS CONSIDERED TO BE RELEVANT Category* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. 1,6,7 2-5, 8-15 X US 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) Local the entire document. ** See the entire document. ** See the entire document. ** See the entire document. ** Social categories of sted document. ** document defining the general state of the art which is not considered to stee or of particular relevance, the claimed invention cannot be supported state claims of support and state of the support of stee claims of support and state of the stee of support stee claims of support stee states of the stee of support stee states of the stee of support stee states of the stee of support stee states of supp	Documentat	ion searched other than minimum documentation to the	extent that such documents are included	in the fields searched				
Category* Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Y See the entire document. Los 5,153,900 A (Nomikos et al) 06 October 1992 (06/10/92) Los 5,153,900 A (Nomikos et al) 06 October 1992 (06/10/92) Los 5,153,900 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) Los 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) Los 6,7 Los 6 the entire document. Purchased ocument defining the general state of the art which is not considered to be particular relevance Los 6 particular relevance Los 6 particular relevance of particular relevance Los 6 particular relevance of particular relevance Los 6 particular relevance, the claimed invention cannot be considered to the expectate area of a publication date of another citation or other special reason (as specified) Los 6 particular relevance, the claimed invention cannot be considered to the content which may throw doubts on pricing date and not in conflict with the applications but cited to understand the principle of or exemit to enough the invention cannot be considered to the content as factor Lose document published principle of or exemit to enough the miremational filing date or priority date and not in conflict with the applications but cited to understand the principle of or exemit to enough the invention cannot be considered to the content in a factor Lose document in particular relevance; the claimed invention cannot be considered to the content in a factor Lose document in particular relevance; the claimed invention cannot be considered to the content in a factor Lose document in particular relevance; the claimed invention cannot be considered to the content in a factor Lose document in particular relevance; the claimed invention cannot be considered to the content in a factor Lose document in particular relevance; the claimed invention cannot be considered to the content in a factor Lose document in the content in a factor Lose of the content in a factor Lose of the content in a factor Lose o	Electronic d	lata base consulted during the international search (na	ame of data base and, where practicable	, search terms used)				
US 5,153,900 A (Nomikos et al) 06 October 1992 (06/10/92) Li,6,7 See the entire document. US 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) See the entire document. See patent family annex. The document defining the general tast of the art which is not considered to be of particular relevance the elained invention cannot be considered for order or other special reason (as specified) document published prior to the international filing date but later than the document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the international filing date but later than document published prior to the i	C. DOC	UMENTS CONSIDERED TO BE RELEVANT						
See the entire document. See the entire document See the entire Se	Category*	Citation of document, with indication, where ap	propriate, of the relevant passages	Relevant to claim No.				
A US 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96) Further documents are listed in the continuation of Box C. See the entire documents. Special categories of cited documents: As document defining the general state of the art which is not considered to be of particular relevances error earlier document published on or after the international filing date of countent to the international filing date or priority date and not un conflict with the application but cited to understand the principle of theory underlying the invention cannot be considered to establish the publication date of another citation or other special reason (as specified) "O' document referring to an oral disclosure, use, exhibition or other special featon (as specified) "O' document referring to an oral disclosure, use, exhibition or other special reason (as specified) "O' document published prior to the international filing date but later than the principly date claimed international search the principle date claimed international search the principle date claimed international search to provide the publication date of another citation or other special reason (as specified) To JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231	Y							
Further documents are listed in the continuation of Box C. See patent family annex. * Special categories of cited documents: *A* document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance to be of particular relevance to the of particular relevance to the serior document which may throw doubte on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) *O* document which may throw doubte on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) *O* document referring to an oral disclosure, use, exhibition or other means *P* document published prior to the international filing date but later than the priority date claimed *A* document published prior to the international filing date but later than the priority date claimed *A* document metering to an oral disclosure, use, exhibition or other means *P* document published prior to the international filing date but later than the priority date claimed *A* document metering to an oral disclosure, use, exhibition or other means *Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231	Α			*				
Further documents are listed in the continuation of Box C. See patent family annex. Special categories of cited documents: document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance arrived document updished and or of or the principle or theory underlying the invention cannot be considered to be of particular relevance to the catablish the publication date of another citation or other special reason (as specified) document which may throw doubte on priority claim(c) or which is cited to establish the publication date of another citation or other special reason (as specified) document published prior to the international filing date but later than the priority date claimed "P" document published prior to the international filing date but later than the priority date claimed Ty JULY 2000 Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231	X	US 5,576,549 A (Hell et al) 19 NOVEMBER 1996 (19/11/96)		1,6,7				
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance "E" earlier document published on or after the international filing date document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "&" document member of the same patent family Date of mailing of the international search report 0 2 AUG 2000 Authorized officer Craig Church Authorized officer Craig Church	A	See the entire document.		2-5,8-15				
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance "E" earlier document published on or after the international filing date document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "&" document member of the same patent family Date of mailing of the international search report 0 2 AUG 2000 Authorized officer Craig Church Authorized officer Craig Church								
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance; the claimed invention cannot be considered novel or cannot be considered novel or cannot be considered novel or cannot be considered to involve an inventive step when the document is staten alone to septimize the priority date claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention cannot be considered novel or cannot be considered novel o								
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance; the claimed invention cannot be considered novel or cannot be considered novel or cannot be considered novel or cannot be considered to involve an inventive step when the document is staten alone to septimize the priority date claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention cannot be considered novel or cannot be considered novel o								
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance; the claimed invention cannot be considered novel or cannot be considered novel or cannot be considered novel or cannot be considered to involve an inventive step when the document is staten alone to septimize the priority date claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention cannot be considered novel or cannot be considered novel o								
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance; the claimed invention cannot be considered novel or cannot be considered novel or cannot be considered novel or cannot be considered to involve an inventive step when the document is staten alone to septimize the priority date claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention cannot be considered novel or cannot be considered novel o								
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance; the claimed invention cannot be considered novel or cannot be considered novel or cannot be considered novel or cannot be considered to involve an inventive step when the document is staten alone to septimize the priority date claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention cannot be considered novel or cannot be considered novel o		_	į					
Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance to be of particular relevance "E" earlier document published on or after the international filing date document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art "&" document member of the same patent family Date of mailing of the international search report 0 2 AUG 2000 Authorized officer Craig Church Authorized officer Craig Church	Further documents are listed in the continuation of Box C. See potent family annex							
"E" earlier document published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed "Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "A" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of mailing of the international search report 0 2 AUG 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231	• Sp	pecial categories of cited documents:	<u> </u>	ernational filing date or priority				
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 considered novel or cannot be considered to involve an inventive step when the document is taken alone considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of mailing of the international search report 0 2 AUG 2000 Authorized officer Craig Church Cannot be considered to involve an inventive step when the document is combined invention cannot be considered to involve an inventive step when the document is taken alone considered to involve an inventive step when the document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document is taken alone considered to involve an inventive step when the document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document is taken alone considered to involve an inventive step when the document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document is considered to involve an inventive step when the document of particular relevance, the claimed invention cannot be considered to involve an inventive step when the document is document of particular relevance, the claimed invention cannot be considered t								
cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of mailing of the international search report 0 2 AUG 2000 Authorized officer Craig Church Craig Church Occument of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family Date of mailing of the international search report Caraig Church O 2 AUG 2000			considered novel or cannot be conside	e claimed invention cannot be red to involve an inventive step				
document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Considered to involve an inventive step when the document is considered to involve an inventive step and inventive step and involve an	cit	ted to establish the publication date of another citation or other	"Y" document of particular relevance; the	e claimed invention cannot be				
Date of the actual completion of the international search 17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Date of mailing of the international search report 0 2 AUG 2000 Authorized officer Craig Church	m		combined with one or more other suc	h documents, such combination				
17 JULY 2000 Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Authorized officer Craig Church			"&" document member of the same paten	t family				
Name and mailing address of the ISA/US Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Authorized officer Craig Church	Date of the	actual completion of the international search						
Commissioner of Patents and Trademarks Box PCT Washington, D.C. 20231 Craig Church	17 JULY	2000	0 2 AUG 20	UU				
Washington, D.C. 20231	Commission		1 / marga					
	Washingto		77/04/0	7				